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(54) Title: METHOD OF INHIBITING RECEPTOR TYROSINE KINASES WITH AN EXTRACELLULAR ANTAGONIST
AND AN INTRACELLULAR ANTAGONIST

(57) Abstract: The present invention relates to methods of inhibiting receptor tyrosine kinases by utilizing a combination of both an extracellular and an intracellular RTK antagonist. The extracellular RTK antagonist is a biological molecule or a small molecule that inhibits activation of the receptor tyrosine kinase by interacting with the extracellular binding region of the receptor. The intracellular RTK antagonist is a biological molecule or small molecule that inhibits tyrosine kinase activity of the receptor tyrosine kinase by interacting with the receptor's intracellular region bearing a kinase domain or by interacting with an intracellular protein involved in the signaling pathway of the receptor tyrosine kinase. The present invention also provides methods of treating tyrosine kinase-dependent diseases, and compositions for use in such methods thereof, by administering a combination of both an extracellular and an intracellular RTK antagonist.

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